

AMENDMENTS TO THE CLAIMS:

This listing of claims will replace all prior versions, and listings, of claims in the application:

1. (currently amended) A pharmaceutical composition comprising a mixture of:

- (a) ~~an active macromolecular principle~~ a polypeptide or protein;
- (b) a non-conjugated bile acid or salt; and
- (c) an additive chosen from
 - (i) propyl gallate or a linear or branched chain C₁₋₁₂ alkyl, C₁₋₁₂ alkyloxy, C₁₋₁₂ alkylthio or C₂₋₁₂ alkenyl ester of gallic acid which is optionally substituted with one or more groups which are the same or different and are selected from halogen and a linear or branched chain C₁₋₁₂ alkyl, C₁₋₁₂ alkyloxy, C₁₋₁₂ alkylthio or C₂₋₁₂ alkenyl ester;
 - (ii) butyl hydroxy anisole, or butyl hydroxy anisole wherein the methyl group or the methoxy group linked to the aromatic ring and/or the hydrogen *ortho* to the hydroxyl group is/are replaced by one or more groups which are the same or different and are selected from linear or branched chain C₁₋₁₂ alkyl, C₁₋₁₂ alkyloxy, C₁₋₁₂ alkylthio and C₂₋₁₂ alkenyl, either unsubstituted or substituted in any position by one or more halogen atoms; and
 - (iii) a mixture of (i) and (ii)

wherein the mixture comprises at least 1% by weight of the additive (c) and the composition is coated with an enteric coating which becomes permeable at a pH from 3 to 7.

2. (original) A composition according to claim 1, which comprises less than 5% by weight of water.

3.-4 (Canceled).

5. (currently amended) A composition according to claim 1, wherein the ratio by weight of the non-conjugated bile salt+additive (b + c) to ~~active macromolecular principle~~the polypeptide or protein is at least 5:1.

6. (previously presented) A composition according to claim 1, wherein the mixture is in the form of a solution or a microparticulate dispersion.

7. (previously presented) A composition according to claim 1, wherein the mixture is in solid form.

8. (Canceled).

9. (currently amended) A composition according to claim 8, where the ~~active macromolecular principle~~polypeptide or protein is chosen from insulin, calcitonin, growth hormone, parathyroid hormone, ~~or~~ erythropoietin, GLP1 and GCSF, and derivatives and analogues thereof, either synthetic or from natural sources, conforming to structures derived from either human or animal origin.

10. (currently amended) A composition according to claim 9, where the ~~active macromolecular principle~~polypeptide or protein is insulin, calcitonin, parathyroid hormone or a derivative or

analogue thereof, either synthetic or from natural sources, conforming to structures derived from either human or animal origin.

11. (currently amended) A composition according to claim 10, wherein the ~~active~~
~~macromolecular principle~~ polypeptide or protein is insulin or a derivative or analogue thereof,
either synthetic or from natural sources, conforming to structures derived from either human or
animal origin, and the composition further comprises an insulin sensitizing agent.

12. (currently amended) A composition according to claim 1, wherein the ~~non-conjugated bile~~
~~acid or salt~~ component (b) is chenodeoxycholate.

13. (previously presented) A composition according to claim 1, wherein the additive is chosen
from propyl gallate or an analogue or a derivative thereof, including esters of gallic acid, where
the esters may be linear or branched chain C₁₋₁₂ alkyl, C₁₋₁₂ alkyloxy, C₁₋₁₂ alkylthio or C₂₋₁₂
alkenyl esters, and the compounds are optionally substituted with halogen, linear or branched
chain C₁₋₁₂ alkyl, C₁₋₁₂ alkyloxy, C₁₋₁₂ alkylthio or C₂₋₁₂ alkenyl esters.

14. (previously presented) A composition according to claim 1, wherein the additive is chosen
from BHA or an analogue or derivative thereof, including analogues and derivatives of hydroxy
anisole where the methyl group or the methoxy group linked to the aromatic ring and/or the
hydrogen ortho to the hydroxyl group are replaced by linear or branched chain C₁₋₁₂ alkyl, C₁₋₁₂
alkyloxy, C₁₋₁₂ alkylthio or C₂₋₁₂ alkenyl, either unsubstituted or substituted in any position,
especially by halogen atoms.

15.-18. (Canceled).

19. (currently amended) A method according to claim ~~18~~26 wherein the ~~molecule(s)/active macromolecular principle~~polypeptide or protein to be absorbed is chosen ~~from~~from insulin, calcitonin, growth hormone, parathyroid hormone, ~~or~~ erythropoietin, GLPI and GCSF, and derivatives and analogues thereof, either synthetic or from natural sources, conforming to structures derived from either human or animal origin.

20. (currently amended) A method according to claim 19, wherein the ~~molecule(s)/active macromolecular principle~~polypeptide or protein to be absorbed is insulin, calcitonin, parathyroid hormone or a ~~derivatives~~derivative or analogue thereof, either synthetic or from natural sources, conforming to structures derived from either human or animal origin.

21. (currently amended) A method according to claim 20, wherein the ~~molecule(s)/active macromolecular principle~~polypeptide or protein to be absorbed is insulin or a ~~derivatives~~derivative or analogue thereof, either synthetic or from natural sources, conforming to structures derived from either human or animal origin, and an insulin sensitizing agent is also present.

22. (previously presented) A method according to claim 26, wherein the composition comprises less than 5% by weight of water.

23. (currently amended) A method according to claim 26, ~~which comprises incorporating wherein~~
~~the active macromolecular principle (s) to be absorbed into the aromatic alcohol in the form~~
~~of polypeptide or protein, the non-conjugated bile acid or salt and the additive are formulated as a~~
solution, ~~as a microparticulate dispersion or as a solid.~~

24. (currently amended) A method of enhancing the absorption of ~~an active macromolecular~~
~~principle~~ a polypeptide or protein in a patient, which method comprises administering to said
patient a composition as defined in claim 1.

25. (Canceled).

26. (currently amended) A method of enhancing the absorption of ~~macromolecules~~ polypeptides
or proteins across the intestinal wall in a human or animal body, which method comprises
administering a non-conjugated bile acid or salt, together with an additive chosen from:

- (i) propyl gallate or a linear or branched chain C₁₋₁₂ alkyl, C₁₋₁₂ alkyloxy, C₁₋₁₂
alkylthio or C₂₋₁₂ alkenyl ester of gallic acid which is optionally substituted with
one or more groups which are the same or different and are selected from halogen
and a linear or branched chain C₁₋₁₂ alkyl, C₁₋₁₂ alkyloxy, C₁₋₁₂ alkylthio or C₂₋₁₂
alkenyl ester;
- (ii) butyl hydroxy anisole, or butyl hydroxy anisole wherein the methyl group or the
methoxy group linked to the aromatic ring and/or the hydrogen *ortho* to the
hydroxyl group is/are replaced by one or more groups which are the same or
different and are selected from linear or branched chain C₁₋₁₂ alkyl, C₁₋₁₂ alkyloxy,

C₁₋₁₂ alkylthio and C₂₋₁₂ alkenyl, either unsubstituted or substituted in any position
by one or more halogen atoms; and

(iii) a mixture of (i) and (ii)

together with the polypeptide or protein in a pharmaceutical composition, wherein the additive accounts for at least 1% by weight of the total weight of (a) the polypeptide or protein, (b) the non-conjugated bile acid or salt, plus (c) the additive, and wherein the composition is coated with an enteric coating which becomes permeable at a pH from 3 to 7, and which method enhances the absorption of the polypeptides or proteins due to the additive improving the solubility of the bile salt.

27. (new) A pharmaceutical composition according to claim 1, wherein the enteric coating becomes permeable at a pH from 5.5 to 7.

28. (new) A pharmaceutical composition according to claim 27, wherein the enteric coating becomes permeable at a pH from 5.5 to 6.5.

29. (new) A method according to claim 26, wherein the enteric coating becomes permeable at a pH from 5.5 to 7.

30. (new) A method according to claim 29, wherein the enteric coating becomes permeable at a pH from 5.5 to 6.5.